

U.S.S.N. 09/030,571
Cantor et al.
PRELIMINARY AMENDMENT

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ceramics, ~~metals~~, resins, gels, membranes and chips].

Please add claims 65-110 as follows.

- 65. The method of claim 5, wherein the solid support is selected from the group consisting of plastics, ceramics, metals, resin, gels, membranes and chips.—
- 66. The method of claim 5, wherein the solid support is a two-dimensional or a three-dimensional matrix with multiple probe binding sites.—
- 67. The method of claim 1, wherein the probes are labelled with a detectable label.—
- 68. The method of claim 1, wherein the detectable label is selected from the group consisting of a radioisotope, a stable isotope, an enzyme, an antibody, a fluorescent chemical, a luminescent chemical, a chromatic chemical, and a metal.—
- 69. The method of claim 1, wherein the nucleic acids are DNA, RNA, PNA or a combination thereof.—
- sub D3*
C3 —70. An array of nucleic acid probes, wherein each probe has a double-stranded portion, a single stranded portion, and a random nucleotide sequence within the single-stranded portion.—
- 71. The array of claim 70 comprising 4^R different nucleic acid probes, wherein R is the length of a random nucleotide sequence within the single-stranded portion of said probe.—
- 72. The array of claim 70, wherein the double-stranded portion is between about 3-20 nucleotides and the single-stranded portion is between about 3-20 nucleotides.—
- 73. The array of claim 70, wherein the double-stranded portion is between 3-20 nucleotides and the single-stranded portion is between 3-20 nucleotides.—

- Sub F3
- 74. The array of claim 70, wherein the nucleic acid probes are fixed to a solid support.—
- 75. The array of claim 74, wherein the solid support is selected from the group consisting of plastics, ceramics, metals, resins, gels, membranes and chips.—
- 76. The array of claim 74, wherein the solid support is a two-dimensional or a three-dimensional matrix with multiple probe binding sites.—
- 77. The array of claim 70, wherein the probes are labelled with a detectable label.—
- 78. The array of claim 77, wherein the detectable label is selected from the group consisting of a radioisotope, a stable isotope, an enzyme, an antibody, a fluorescent chemical, a luminescent chemical, a chromatic chemical, and a metal.—
- 79. The array of claim 70, wherein the nucleic acids are DNA, RNA, Protein Nucleic Acid (PNA), or a combination thereof.—
- 83 with sub D4
- 80. A method for detecting a target nucleic acid in a biological sample comprising:
- a) contacting the array of probes with the sample, wherein each probe has a double-stranded portion, a single-stranded portion, and a random sequence within the single-stranded portion; and
 - b) identifying hybrids, whereby the target nucleic acid is detected.—
- 81. The method of claim 80, wherein the biological sample is selected from the group consisting of samples of animal tissue, environmental substances, manufacturing products and by-products.—
- 82. The method of claim 81, wherein the animal tissue is obtained from a human.—
- 83. The method of claim 80, further comprising the step of purifying the target nucleic acids detected.—
- 84. The method of claim 80, wherein the set of nucleic acid probes is fixed to
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a solid support.—

- 85. The method of claim 84, wherein the solid support is selected from the group consisting of plastics, ceramics, metals, resin, gels, membranes and chips.—
- 86. The method of claim 84, wherein the solid support is a two-dimensional or a three-dimensional matrix with multiple probe binding sites.—
- 87. The method of claim 80, wherein the target nucleic acids or the probes are labelled with a detectable label.—
- 88. The method of claim 87, wherein the detectable label is selected from the group consisting of radioisotope, a stable isotope, an enzyme, an antibody, a fluorescent chemical, a luminescent chemical, a chromatic chemical, and a metal.—
- Sub D5* —89. A solid support, comprising an array of nucleic acid probes, wherein each probe has a double-stranded portion, a single-stranded portion, and a random sequence within the single-stranded portion.—
- Q3* —90. The solid support of claim 89, wherein the solid support is selected from the group consisting of plastics, ceramics, metals, resin, gels, membranes and chips.—
- 91. The solid support of claim 89, wherein the solid support is a two-dimensional or a three-dimensional matrix with multiple probe binding sites.—
- Sub F4* —92. The solid support of claim 89, wherein the probes are labelled with a detectable label.—
- 93. The solid support of claim 92, wherein the detectable label is selected from the group consisting of radioisotope, a stable isotope, an enzyme, an antibody, a fluorescent chemical, a luminescent chemical, a chromatic chemical, and a metal.—
- 94. The solid support of claim 89, wherein the nucleic acids are DNA, RNA, Protein Nucleic Acid (PNA), or a combination thereof.—

- C3*
- at*
- 95. A method of sequencing a target nucleic acid, comprising the steps of:
- i) hybridizing the target nucleic acid to an array of nucleic acid probes; and
 - ii) determining a hybridization pattern; whereby the target nucleic acid is sequenced by analyzing the hybridization pattern, wherein:
 - a) the nucleic acid target is at least partly single-stranded; and
 - b) each probe comprises a double-stranded portion, a single stranded portion, and a random sequence within the single-stranded portion.—
- 96. The method of claim 95, further comprising the step of ligating the hybridized target to the probe.—
- 97. The method of claim 95, further comprising the step of enzymatically extending a strand of the probe using the hybridized target as a template.—
- 98. The method of claim 97, wherein the probe is enzymatically extended by a DNA polymerase.—
- 99. The method of claim 97, wherein the probe is enzymatically extended by DNA polymerase using a single deoxynucleotide triphosphate or dideoxynucleotide triphosphate.—
- 100. The method of claim 95, further comprising the steps of ligating the hybridized target to the probe, and enzymatically extending a strand of the probe using the hybridized target as a template.—
- 101. The method of claim 100, wherein the probe is enzymatically extended by a DNA polymerase.—
- 102. The method of claim 100, wherein the probe is enzymatically extended by DNA polymerase using a single deoxynucleotide triphosphate or dideoxynucleotide triphosphate.—

- 103. The method of claim 95, wherein the nucleic acids are DNA, RNA, PNA, or a combination thereof.—
- 104. The method of claim 95, wherein the set of nucleic acid probes is fixed to a solid support.—
- 105. The method of claim 104, wherein the solid support is selected from the group consisting of plastics, ceramics, metals, resin, gels, membranes and chips.—
- 106. The method of claim 104, wherein the solid support is a two-dimensional or a three-dimensional matrix with multiple probe binding sites.—
- 107. The method of claim 95, wherein the target nucleic acid or the probes are labelled with a detectable label.—
- 108. The method of claim 107, wherein the detectable label is selected from the group consisting of radioisotope, a stable isotope, an enzyme, an antibody, a fluorescent chemical, a luminescent chemical, a chromatic chemical, and a metal.—
- 109. The method of claim 95, wherein the hybridization pattern is analyzed by a computer.—
- 110. A method of sequencing a target nucleic acid comprising the steps of:
- i) hybridizing the target nucleic acid to an array of nucleic acid probes; and
 - ii) detecting the hybridized target nucleic acid; whereby the target nucleic acid is sequenced, wherein:
 - a) the nucleic acid target is at least partly single-stranded; and
 - b) each probe comprises a double-stranded portion, a single stranded portion, and a random sequence within the single-stranded portion.—